## **WEST Search History**

Hide Items Restore Clear Cancel

DATE: Friday, January 21, 2005

Hide? Set Name Query			Hit Count
$DB=USPT,PGPB,JPAB,DWPI;\ PLUR=YES;\ OP=ADJ$			
	L10	L9 and @pd > 20040624	72
	L9	(L7 or L5) not L3	316
	L8	L7 or L5 not L3	327
	L7	L6 and (heterolog\$ express\$ or express\$ casset\$)	292
	L6	L2 and ischem\$	3412
	L5	L2 and tumor infiltr\$ and (heterolog\$ express\$ or express\$ casset\$)	59
	L4	L2 and (tumor infiltr\$ or heterolog\$ express\$ or express\$ casset\$)	1410
	L3	L2 and (HRE or HIF1-alpha)	117
	L2	L1 and hypox\$	9723
	L1	phagocyte\$ or macrophage\$ or monocyte\$	50484

END OF SEARCH HISTORY

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Virology
LA English
                                                                                                                                                 => FIL BIOSIS EMBASE CAPLUS
LA English

AB Somatic cell hybrid clones between either C57BL/6 or Balb/c mouse
peritoneal ***macrophages*** and two different simian virus 40 (SV40)
***transformed*** human cell lines deficient in ***hypoxanthine***
phosphoribosyltransferase (EC 2.4.2.8; IMP:pyrophosphate
phosphoribosyltransferase) were obtained in ***hypoxanthine***
aminoptenin thymidine selective medium. All the hybrid cell clones
                                                                                                                                                COST IN U.S. DOLLARS
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                                                                                                                                                                                                                 SESSION
                                                                                                                                                                                                   ENTRY
                                                                                                                                                 FULL ESTIMATED COST
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                                                                                                                                                 FILE 'BIOSIS' ENTERED AT 16:45:32 ON 30 NOV 2004
                                                                                                                                                 Copyright (c) 2004 The Thomson Corporatio
     contained the human chromosome 7, which carries the SV40 genome, and were SV40 tumor (T) antigen positive. No hybrid cell clones studied displayed the density dependent inhibition of cell growth characteristic of normal
                                                                                                                                                 FILE 'EMBASE' ENTERED AT 16:45:32 ON 30 NOV 2004
                                                                                                                                                 COPYRIGHT (C) 2004 Elsevier Inc. All rights reserved.
    cells; all clones had a high saturation density and gave origin to cell colonies when plated in soft agar. Since the ***expression*** of the ***transformed*** phenotype was always associated with the presence of the human chromosome 7, which carries the SV40 genome, it is concluded that this chromosome contais gene(s) [Tr gene(s)] coding for ' ***transforming*** factor(s)'.
                                                                                                                                                FILE 'CAPLUS' ENTERED AT 16:45:32 ON 30 NOV 2004
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
                                                                                                                                                 COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)
                                                                                                                                                => s tumor? or cancer?
L1 2777028 TUMOR? OR CANCER?
                                                                                                                                                => s I1 and gene therapy
L2 31230 L1 AND GENE THERAPY
---Logging off of STN---
                                                                                                                                                 => s I2 and macrophages or monocyt? or phagocyt?
L3 311738 L2 AND MACROPHAGES OR MONOCYT? OR PHAGOCYT?
                                                                                                                                                 => s I2 and (macrophages or monocyt? or phagocyt?)
L4 942 L2 AND (MACROPHAGES OR MONOCYT? OR PHAGOCYT?)
Executing the logoff script...
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L5 23755 THERAPEU? (3A) GENE?
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L6 151 L4 AND L5
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2 FILES SEARCHED..
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STN INTERNATIONAL LOGOFF AT 16:28:48 ON 27 FEB 2003
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L8 4 DUP REM L7 (0 DUPLICATES REMOVED)
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                                                                                                                                                 L8 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN
Welcome to STN International Enter x:x
                                                                                                                                                 AN 2000:271946 CAPLUS
LOGINID:ssspta1633cxq
                                                                                                                                                 DN 132:307254
                                                                                                                                                 TI Antigen-binding sites of antibody molecules specific for ***cancer***
PASSWORD:
                                                                                                                                                     antigens
TERMINAL (ENTER 1, 2, 3, OR ?):2
                                                                                                                                                 IN Ring, David B.
                                                                                                                                                PA Chiron Corporation, USA
SO U.S., 57 pp., Cont.-in-part of U.S. 5,629,197.
CODEN: USXXAM
******* Welcome to STN International
                                                                                                                                                 DT Patent
                       Web Page URLs for STN Seminar Schedule - N. America "Ask CAS" for self-help around the clock
                                                                                                                                                LA English
FAN.CNT 2
 NEWS 1
 NEWS 2
 NEWS 3 SEP 01 INPADOC: New family current-awareness alert (SDI) available
                                                                                                                                                     PATENT NO.
                                                                                                                                                                                   KIND DATE
                                                                                                                                                                                                             APPLICATION NO.
                                                                                                                                                                                                                                                  DATE
NEWS 4 SEP 01 New pricing for the Save Answers for SciFinder Wizard within STN Express with Discover!
                                                                                                                                                     US 6054561
                                                                                                                                                                                          20000425 US 1995-483749
                                                                                                                                                                                                                                               19950607
NEWS 5 SEP 01 New display format, HITSTR, available in WPIDSWPINDEXWPIX
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19890425 CA 1985-472301
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19850117 <--
                                                                                                                                                     US 4753894
                                                                                                                                                     CA 1253090
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NEWS 6 SEP 27 STANDARDS will no longer be available on STN NEWS 7 SEP 27 SWETSCAN will no longer be available on STN NEWS 8 OCT 28 KOREAPAT now available on STN
                                                                                                                                                     ZA 8500980
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19961112 JP 1996-81684
 NEWS 9 NOV 18 Current-awareness alerts, saved answer s
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                 search transcripts to be affected by CERAB, COMPUAB, ELCOM, and SOLIDSTATE reloads
                                                                                                                                                     JP 08295700
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2 19850111
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US 1986-842476
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B1
 NEWS 10 NOV 30 PHAR reloaded with additional data
                                                                                                                                                                                            19860321
 NEWS EXPRESS OCTOBER 29 CURRENT WINDOWS VERSION IS V7.01A,
                                                                                                                                                     US 1988-190778
                                                                                                                                                                                             19880508
                                                                                                                                                     US 1994-288981
EP 1985-300877
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CURRENT
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               MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
               AND CURRENT DISCOVER FILÈ IS DATED 11 AUGUST 2004
                                                                                                                                                     JP 1992-95610
                                                                                                                                                                                   АЗ
                                                                                                                                                                                          19920415
NEWS HOURS STN Operating Hours Plus Help Desk Availability NEWS INTER General Internet Information
                                                                                                                                                AB Novel compns, are provided that are derived from antigen-binding sites of lgs having affinity for ***cancer*** antigens. The compns. exhibit
                                                                                                                                                    Igs having affinity for ***cancer*** antigens. The compns. exhibit immunol. binding properties of antibody mols. capable of binding specifically to a human ***bumor*** cell expressing an antigen selected from the group consisting of high mol. wt. mucins bound by 2G3 and 369F10, c-erbB-2 ****tumor*** antigen, an approx. 42 kD glycoprotein, an approx. 55 kD glycoprotein, and the approx. 40, 60, 100 and 200 kD antigens bound by 113F1. A no. of synthetic mols. are provided that include CDR and FR regions derived from same or different to gradients. Also provided are signed so his producer to the processing the control of the provided are signed so his progressing the provided and signed so his progressing the control of the provided are signed so his progressing the provided and signed so have provided that include CDR and FR regions derived from same or different to
 NEWS LOGIN Welcome Banner and News Items
NEWS PHONE Direct Dial and Telecommunication Network Access to STN
 NEWS WWW
                          CAS World Wide Web Site (general information)
Enter NEWS followed by the item number or name to see news on that
                                                                                                                                                     moieties. Also provided are single chain polypeptides wherein VH and VL domains are attached by a single polypeptide linker. The sFv mols. can include ancillary polypeptide moieties which can be bioactive, or which
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  agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation
  of commercial gateways or other similar uses is prohibited and may
                                                                                                                                                     provide a site of attachment for other useful moieties. The compns. are
                                                                                                                                                    useful in specific binding assays, affinity purifin. schemes, drug or toxin targeting, imaging, and ""genetic" or immunol. ""therapeutics" for various ""cancers". The invention thus provides novel polypeptides, the DNAs encoding those polypeptides, expression cassettes
  result in loss of user privileges and other penalties.
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FS 022 Human Genetics

FILE 'HOME' ENTERED AT 16:45:17 ON 30 NOV 2004

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SO PCT Int. Appl., 53 pp. CODEN: PIXXD2
polypeptides.
RE.CNT 44 THERE ARE 44 CITED REFERENCES AVAILABLE FOR THIS
                                                                                                                                                              DT Patent
RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
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L8 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN
                                                                                                                                                                  PATENT NO.
AN 1996:456098 CAPLUS
                                                                                                                                                                      WO 9532282 A1 19951130 WO 1995-US6094 19950517 <-
W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI,
GB, GE, HU, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG,
MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT, UA,
DN 125:107063
                                                                                                                                                              PI WO 9532282
TI Cationic amphiphiles and plasmids for intracellular delivery of
     therapeutic molecules
IN Siegel, Craig S.; Harris, David J.; Lee, Edward R.; Hubbard, Shirley C.
     Cheng, Seng H.; Eastman, Simon J.; Marshall, John; Scheule, Ronald K.;
                                                                                                                                                                      RW: KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE,
     Yew, Nelson S.; et al.
       Genzyme Corporation, USA
                                                                                                                                                                           SN, TD, TG
SO PCT Int. Appl., 152 pp.
CODEN: PIXXD2
                                                                                                                                                                                                         19970624 US 1994-246427
19951218 AU 1995-25515
19961118 ZA 1995-4027
                                                                                                                                                                  US 5641657
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DT Patent
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LA English
FAN.CNT 11
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                                                                  APPLICATION NO.
    PATENT NO.
                                    KIND DATE
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                                                                                                                                                                                                  T2 19980127 JP 1995-530365
A2 20030218 JP 2002-142609
                                                                                                                                                                   JP 10500850
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PI WO 9618372
                                      A2 19960620 WO 1995-US16174
A3 19960906
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                                                                                                                                                                   US 5958400
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     WO 9618372
                                     АЗ
        W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ,
                                                                                                                                                                                                   A 19940519
A3 19950517
W 19950517
                                                                                                                                                              PRAI US 1994-246427
                                                                                                                                                                   JP 1995-530365
                                                                                                                                                                   WO 1995-US6094
                                                                                                                                                              AB Human interleukin 6 splice variant (IL-6SV) polypeptides and DNA (or RNA) encoding such polypeptides and a procedure for producing such polypeptides by recombinant techniques is disclosed. Also disclosed are methods for
        RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR,
     NE, SN, TD, TG
US 5650096 A
                                                                                                                                                                  utilizing such polypeptides for identifying antagonists and agonists to such polypeptides and methods of using the polypeptide and antagonists
                                            19970722
                                                               US 1994-352479
                                                                                                      19941209
                                                                                                                                                                  therapeutically to treat platelet reducing conditions, shock syndromes, as an antiviral agent, to inhibit proliferation of leukemic cells, to improve
     US 5747471
                                            19980505
                                                                US 1995-540867
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     US 6071890
AU 9645161
                                    A
A1
                                           20000606
19960703
                                                                US 1995-545473
                                                                                                      19951019
                                                                AU 1996-45161
                                                                                                       19951208 <--
                                                                                                                                                                  the toxic activity of human lymphocytes for killing ***cancer*
                                                                                                                                                                  cells, for use in cell transplant therapy, and inflammation. Also disclosed are diagnostic methods for detecting a mutation in the IL-6SV
     AU 716706
                                           20000302
     EP 799059
                                  A1
B1
                                           19971008
                                                               EP 1995-943769
                                                                                                     19951208
                                                                                                                                                                  discissed are diagnostic methods for detecting a mutation in the IL-6SV nucleic acid sequences and detecting a level of the polypeptide in a sample derived from a host. The 507-bp cDNA coding for the 167-amino-acid, mature form of IL-6SV was isolated and sequenced from a cDNA library derived from activated human ""macrophages". The
     EP 799059
                                           20020731
     R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE
JP 10510813 T2 19981020 JP 1995-519236 19951208
                                        20020815 AT 1995-943769
A 19980402 CA 1997-2268945
I 19980417 AU 1997-32315
                                  Ε
     AT 221390
                                                                                                    19951208
                                                                                                                                                                  cDNA library derived from activated human ***macrophages***. The examples described the bacterial expression of IL-6SV using the expression vector pQE-60 with a His6 tag and purifn. of the protein from recombinant Escherichia coli on a Ni-Chelate column. Expression via ***gene*** ***therapy*** is achieved by inserting the CDNA into Moloney murine sarcoma virus-derived vector pMV-7 for the transduction of fibroblasts.
                                   AA
     CA 2268945
AU 9732315
                                                                                                        19970610
                                                                                                       19970610
                                    A1
     AU 736143
                                   B2 20010726
                                           20000614 EP 1997-927989
                                                                                                       19970610
     EP 1007003
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IF FI
     JP 2001500897
                                             20010123 JP 1998-515603
                                                                                                        19970610
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                                       T2
     US 2002013282
                                       A1 20020131 US 1998-166074
                                                                                                                                                              AN 1995:210275 CAPLUS
                                          A 199412
19950926
PRAI US 1994-352479
                                                   19941209
                                                                                                                                                              DN 122:532
     US 1995-4344P
                                                                                                                                                              TI Adenovirus-mediated ***gene*** ***therapy*** of experimental
     US 1995-4399P
                                       Р
                                              19950927
                                                                                                                                                             gliomas
AU Perez-Cruet, M. J.; Trask, T. W.; Chen, S-H.; Goodman, J. C.; Woo, S. L. C.; Grossman, R. G.; Shine, H. D.
CS Dep. Neurosurgery, Baylor Coll. Med., Houston, TX, USA
SO Journal of Neuroscience Research (***1994
     US 1995-540867
                                               19951011
     US 1995-545473
                                               19951019
    WO 1995-US16174
WO 1997-US9748
                                         w
                                                   19951208
                                                19970610
OS MARPAT 125:107063
                                                                                                                                                                  CODEN; JNREDK; ISSN: 0360-4012
AB Novel cationic amphiphiles are provided that facilitate transport of biol. active (therapeutic) mols, into cells. The amphiphiles contain lipophilic
                                                                                                                                                              PB Wiley-Liss
                                                                                                                                                                     Journal
     groups derived from steroids, from mono or dialkylamines, or from alkyl or
                                                                                                                                                                    English
    acyl groups; and cationic groups, protonatable at physiol. pH, derived from amines, alkylamines or polyalkylamines. Thus, N4-spermidine cholesteryl carbamate provided an apprx.20-fold enhancement of the transfection ability of plasmid pCMVHI-CAT (chloramphenicol
                                                                                                                                                                     The efficacy of adenovirus (ADV)-mediated ***gene*** ***therapy**
                                                                                                                                                                  to treat brain ***tumors*** was tested in a syngeneic glioma model.
***Tumor*** cells were transduced in situ with a replication-defective
                                                                                                                                                                  ADV carrying the herpes simplex virus thymidine kinase (HSV-tk) gene
                                                                                                                                                                  controlled by the Rous sarcoma virus promoter. Expression of the HSV-tk gene enables the transduced cell to convert the drug ganciclovir to a form that is cytotoxic to dividing cells. ***Tumors*** were generated in
     acetyltransferase-encoding) in mice. There are provided also therapeutic
     compns. prepd. typically by contacting a dispersion of one or more cationic amphiphiles with the therapeutic mols. Therapeutic mols. that
                                                                                                                                                                  that is cytotoxic to dividing cells.

Fischer 344 rats by stereotaxic implantation of 9L gliosarcoma cells into the caudate nucleus. Eight days later, the ***tumor*** were injected
     can be delivered into cells according to the practice of the invention include DNA, RNA, and polypeptides. Representative uses of the
    include DNA, RNA, and polypeptides. Representative uses of the therapeutic compns, of the invention include providing ""gene""
""therapy"", and delivery of antisense polynucleotides of biol. active polypeptides to cells. With respect to ""therapeutic" compns, for ""gene"" "therapy"", the DNA is provided typically in the form of a plasmid for complexing with the cationic amphiphile. Novel and highly effective plasmid constructs are also disclosed, including those that are particularly effective at providing ""gene"" ""therapy"" for clin. conditions complicated by inflammation. Several
                                                                                                                                                                   either with the ADV carrying the HSV-tk (ADV-tk) gene or a control ADV
                                                                                                                                                                  vector confg the .beta_galactosidase (ADV-tx) gene and the rats were treated with either ganciclovir or saline. ***Turnor*** size was
                                                                                                                                                                  measured 20 days after implantation of 9L cells or at death. Rats treated with ADV- beta gal and ganciclovir or with ADV-tk and saline had large ""tumors"". No ""tumors" were detected in animals treated with
                                                                                                                                                                  ADV-tk and with ganciclovir at doses .gtoreq.80 mg/kg. An infiltrate of ""macrophages*" and lymphocytes at the injection site in animals treated with ADV-tk and ganciclovir indicated an active local immune reaction. In survival studies, all animals treated with ADV-tk and
     vectors were constructed for improved delivery of the gene the cystic
     fibrosis transmembrane conductance regulator (CFTR) under control of the human cytomegalovirus promoter/enhancer during cationic
                                                                                                                                                                  ganciclovir have remained alive longer than 80 and up to 120 days after
     amphiphile-mediated gene transfer. Addnl., targeting of organs for
""gene"" ""therapy"" by i.v. administration of therapeutic
compns. is described. Syntheses are described for N4-spermine cholesteryl
                                                                                                                                                                  ***tumor*** induction whereas all untreated animals died by 22 days.

These results demonstrate that ADV-mediated transfer of HSV-tk to glioma
                                                                                                                                                                  cells in vivo confers sensitivity to ganciclovir, and represents a potential method of treatment of brain ***tumors***
     carbamate, N4-(N'-cholesteryl carbamate glycineamide)-spermine, N4-spermidine-2,3-dilauryloxypropylamine, and N4-spermine-2,3-
     dilauryloxypropylamine.
L8 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN
                                                                                                                                                                  (FILE 'HOME' ENTERED AT 16:45:17 ON 30 NOV 2004)
AN 1996:91928 CAPLUS
DN 124:143605
TI Cloning and ***gene*** ***therapy*** of an interleukin-6 splice
                                                                                                                                                                  FILE 'BIOSIS, EMBASE, CAPLUS' ENTERED AT 16:45:32 ON 30 NOV 2004 2777028 S TUMOR? OR CANCER?
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comprising those DNAs, and methods of inducing the prodn. of the

variant and its therapeutic and diagnostic uses IN Ruben, Steven; Li, Haodong; Adams, Mark D.

PA Human Genome Sciences, Inc., USA

31230 S L1 AND GENE THERAPY

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           942 S L2 AND (MACROPHAGES OR MONOCYT? OR PHAGOCYT?)
23755 S THERAPEU? (3A) GENE?
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NE, SN, TD, TG
               4 DUP REM L7 (0 DUPLICATES REMOVED)
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L9 89870 L1 AND (MACROPHAGES OR MONOCYT? OR PHAGOCYT?)
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2 FILES SEARCHED.
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             22 L10 AND PY<=1996
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2 19981020 JP 1995-519236

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YOU HAVE REQUESTED DATA FROM 18 ANSWERS - CONTINUE? Y/(N):y
                                                                                                                                                                                T2 20010123 JP 1998-515603
A1 20020131 US 1998-166074
A 19941209
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L12 ANSWER 1 OF 18 CAPLUS COPYRIGHT 2004 ACS on STN
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PRAI US 1994-352479
AN 2000:271946 CAPLUS
                                                                                                                                                                                                                                             19981005
DN 132:307254
                                                                                                                                                  US 1995-4344P
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TI Antigen-binding sites of antibody molecules specific for ***cancer***
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                                                                                                                                                                                        19950927
    antigens
                                                                                                                                                  US 1995-540867
IN Ring, David B.
                                                                                                                                                                                        19951011
PA Chiron Corporation, USA
SO U.S., 57 pp., Cont.-in-part of U.S. 5,629,197.
CODEN: USXXAM
                                                                                                                                                  US 1995-545473
WO 1995-US16174
                                                                                                                                                                                 Â<sub>W</sub>
                                                                                                                                                                                        19951019
                                                                                                                                                                                           19951208
                                                                                                                                                  WO 1997-US9748
                                                                                                                                                                                          19970610
                                                                                                                                             OS MARPAT 125:107063
DT Patent
                                                                                                                                             AB Novel cationic amphiphiles are provided that facilitate transport of biol.
LA English
FAN.CNT 2
                                                                                                                                                  active (therapeutic) mols. into cells. The amphiphiles contain lipophilic
                                                                                                                                                  groups derived from steroids, from mono or dialkylamines, or from alkyl or
    PATENT NO.
                                 KIND DATE
                                                            APPLICATION NO.
                                                                                               DATE
                                                                                                                                                  groups derived from sterious, from fronts of diadisynamines, of normality of acyl groups, and cationic groups, protonatable at physiol, pH, derived from amines, alkylamines or polyalkylamines. Thus, N4-spermidine cholesteryl carbamate provided an .apprx.20-fold enhancement of the transfection ability of plasmid pCMVHI-CAT (chloramphenicol acetyltransferase-encoding) in mice. There are provided also therapeutic compns. prepd. typically by contacting a dispersion of one or more
PI US 6054561
                                         20000425
                                                           US 1995-483749
                                                                                             19950607
    US 4753894
                                        19880628
                                                         US 1985-690750
                                                                                           19850111 <--
                                                                                            19850117 <--
    CA 1253090
                                        19890425
                                                         CA 1985-472301
                                 Α1
    ZA 8500980
                                     19861029 ZA 1985-980
19890815 AT 1985-300877
                                                                                        19850208 <--
                                                                                         19850208 <--
    AT 44769
                              Ε
     US 5169774
                                        19921208
                                                        US 1988-190778
                                                                                            19880506 <--
                                                                                                                                                  cationic amphiphiles with the therapeutic mols. Therapeutic mols. that
                                                                                                                                                  can be delivered into cells according to the practice of the invention include DNA, RNA, and polypeptides. Representative uses of the therapeutic compns. of the invention include providing gene therapy, and delivery of antisense polynucleotides of biol. active polypeptides to cells. With respect to ***therapeutic*** compns. for ****gene****
                                       19970513 US 1994-288981
19961112 JP 1996-81684
    US 5629197
                                                                                            19940811
                                                                                           19960403 <--
    JP 08295700
                                 A2
                                           19840208
19850111
PRAI US 1984-577976
                                        B2
    US 1985-690750
                                   A2
    US 1986-842476
                                           19860321
                                   B1
                                           19880508
19940811
                                                                                                                                                  therapy, the DNA is provided typically in the form of a plasmid for complexing with the cationic amphiphile. Novel and highly effective
    US 1988-190778
    US 1994-288981
                                    A2
    EP 1985-300877
                                          19850208
                                                                                                                                                  plasmid constructs are also disclosed, including those that are
                                                                                                                                                  particularly effective at providing gene therapy for clin. conditions complicated by inflammation. Several vectors were constructed for improved delivery of the gene the cystic fibrosis transmembrane conductance regulator (CFTR) under control of the human cytomegalovirus
    JP 1992-95610
                                  АЗ
                                         19920415
AB Novel compns. are provided that are derived from antigen-binding sites of lgs having affinity for ***cancer*** antigens. The compns. exhibit
    Igs having attinity for "cancer" antigens. The compine, swhibit immunol, binding properties of antibody mols, capable of binding specifically to a human ""tumor"" cell expressing an antigen selected from the group consisting of high mol. wt. mucins bound by 2G3 and 369F10, c-erbB-2 ""tumor"" antigen, an approx. 42 kD glycoprotein, an approx. 45 kD glycoprotein, and the approx. 40, 60, 100
                                                                                                                                                  promoter/enhancer during cationic amphiphile-mediated gene transfer.
                                                                                                                                                  Addnl., targeting of organs for gene therapy by i.v. administration of therapeutic compns. is described. Syntheses are described for N4-spermine
                                                                                                                                                  cholesteryl carbamate, N4-(N'-cholesteryl carbamate glycineamide)-
    and 200 kD antigens bound by 113F1. A no. of synthetic mols, are provided that include CDR and FR regions derived from same or different Ig
                                                                                                                                                  spermine, N4-spermidine-2,3-dilauryloxypropylamine, and N4-spermine-2,3-dilauryloxypropylamine.
    moieties. Also provided are single chain polypeptides wherein VH and VL domains are attached by a single polypeptide linker. The sFv mols. can include ancillary polypeptide moieties which can be bioactive, or which
                                                                                                                                             L12 ANSWER 3 OF 18 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL
                                                                                                                                             RIGHTS RESERVED.
                                                                                                                                             on STN
AN 96101196 EMBASE
    provide a site of attachment for other useful moieties. The compns. are
    useful in specific binding assays, affinity purifn. schemes, drug or toxin targeting, imaging, and ***genetic*** or immunol. ***therapeutics*** for various ***cancers***. The invention thus provides novel
                                                                                                                                             DN 1996101196
TI ***Monocyte*** /macrophage activation by immunostimulators: Role in ***cancer*** therapy.
    polypeptides, the DNAs encoding those polypeptides, expression cassettes comprising those DNAs, and methods of inducing the produc of the
                                                                                                                                             AU Hennemann B.; Andereesen R.
CS Abteilung Hamatologie/Onkologie, Klinik/Poliklinik Innere Medizine I,
Klinikum Universitat Regensburg, 93042 Regensburg, Germany
    polypeptides
RE.CNT 44 THERE ARE 44 CITED REFERENCES AVAILABLE FOR THIS
                                                                                                                                             SO Clinical Immunotherapeutics, (1996) 5/4 (294-308). ISSN: 1172-7039 CODEN: CIMMEA
RECORD
            ALL CITATIONS AVAILABLE IN THE RE FORMAT
                                                                                                                                             CY New Zealand
L12 ANSWER 2 OF 18 CAPLUS COPYRIGHT 2004 ACS on STN AN 1996:456098 CAPLUS
                                                                                                                                             DT Journal; General Review
                                                                                                                                             FS 016 Cancer
DN 125:107063
                                                                                                                                                  026 Immunology, Serology and Transplantation
                                                                                                                                                            Clinical Biochemistry
TI Cationic amphiphiles and plasmids for intracellular delivery of
                                                                                                                                                  029
    therapeutic molecules
                                                                                                                                                            Drug Literature Index
    Siegel, Craig S.; Harris, David J.; Lee, Edward R.; Hubbard, Shirley C.; Cheng, Seng H.; Eastman, Simon J.; Marshall, John; Scheule, Ronald K.;
                                                                                                                                             LA English
SL English
                                                                                                                                             AB Cells of the ***monocyte*** /macrophage lineage are considered to be of special importance in host defence against tumour growth. There is
     Yew, Nelson S.; et al.
PA Genzyme Corporation, USA
SO PCT Int. Appl., 152 pp.
CODEN: PIXXD2
                                                                                                                                                  experimental and clinical evidence that in malignant disease the generation of cytotoxic ***macrophages*** is impaired, Both defective cell maturation and loss of responsiveness to activation have been
DT Patent
LA English
                                                                                                                                                  described, Immunotherapeutic strategies to stimulate macrophage tumour
                                                                                                                                                  cytotoxicity make use of activating compounds such as interferon-gamma. (IFN.gamma.), endotoxin (lipopolysaccharide) and other cytokines that are
FAN CNT 11
    PATENT NO.
                                 KIND DATE
                                                            APPLICATION NO.
                                                                                                DATE
                                                                                                                                                  administered systemically. Subcutaneous treatment with low-dose IFN.gamma. given on a weekly schedule achieved an objective response of 4 to 30% in
PI WO 9618372
                                   A2 19960620 WO 1995-US16174
                                                                                                 19951208 <--
```

311738 S L2 AND MACROPHAGES OR MONOCYT? OR PHAGOCYT?

A3 19960906

WO 9618372

were given intravenously and achieved an objective response in 9% and stable disease in 49% of patients with renal cell carcinoma, Lipopolysaccharide given intravenously induced a profound immunological response in the recipient, Antitumour activity was seen in 25% of patients with advanced \*\*\*cancer\*\*\* Adoptive immunotherapy with \*\*\*macrophages\*\*\* generated in vitro is a treatment modality designed to correct for defective in vivo maturation of \*\*\*monocyte\*\*\*. Preclinical data in murine models showed a remarkable antitumour effect of transferred cells. Activated \*\*\*macrophages\*\*\* given locally or via intravenous injection inhibited tumour growth of Lewis lung carcinoma by 30 to 40% in C57B16 mice. Clinical trials with local and systemic transfe of autologous cytotoxic \*\*\*macrophages\*\*\* showed the induction of neopterin, interleukin-6 and thrombin-antithrombin complexes in the recipient. The antitumour activity of local therapy was evident from the disappearance of malignant ascites upon intraperitoneal cell application. However, as reported by several groups, intravenous cell transfer has yielded conflicting results and only minor turnour responses were seen Here, further improvements in culture technique and mode of cell activation are being developed. In addition, \*\*\*macrophages\*\*\* coul
be used as a target of \*\*\*gene\*\*\* transfer experiments. The
\*\*\*therapeutic\*\*\* value of this technique needs careful investigation.

patients with metastatic renal cell carcinoma, Higher doses of IFN.gamma.

L12 ANSWER 4 OF 18 CAPLUS COPYRIGHT 2004 ACS on STN AN 1996:91928 CAPLUS DN 124:143605

TI Cloning and gene therapy of an interleukin-6 splice variant and its therapeutic and diagnostic uses

IN Ruben, Steven; Li, Haodong; Adams, Mark D. PA Human Genome Sciences, Inc., USA

PCT Int. Appl., 53 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

VO 9532282 A1 19951130 WO 1995-US6094 19950517 <-W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, 19950517 <--PI WO 9532282 MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT, UA,

UZ, VN RW: KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

US 5641657 19970624 US 1994-246427 19940519 19951218 AU 1995-25515 19961118 ZA 1995-4027 AU 9525515 19950517 <---ZA 9504027 19950517 <--EP 759980 19970305 EP 1995-919845 19950517 R: BE, CH, DE, FR, GB, LI, NL - 10500850 T2 19980127 JP 1995-530365 19950517

1∠ 19980127 JP 1995-530365 JP 2003047483 A2 20030218 JP 2002-142609 US 5958400 A 19990928 US 1996-766620 PRAI US 1994-246427 A 19940519 JP 1995-530365 19950517 19961213

RAI US 1994-246427 A 19940519 JP 1995-530365 A3 19950517 WO 1995-US6094 W 19950517

AB Human interleukin 6 splice variant (IL-6SV) polypeptides and DNA (or RNA) encoding such polypeptides and a procedure for producing such polypeptides by recombinant techniques is disclosed. Also disclosed are methods for utilizing such polypeptides for identifying antagonists and agonists to such polypeptides and methods of using the polypeptide and antagonists therapeutically to treat platelet reducing conditions, shock syndromes, as an antiviral agent, to inhibit proliferation of leukemic cells, to improve the toxic activity of human lymphocytes for killing \*\*\*cancer\*\*\* cells, for use in cell transplant therapy, and inflammation. Also cells, for use in cell transplant therapy, and inharmation. Also disclosed are diagnostic methods for detecting a mutation in the IL-6SV nucleic acid sequences and detecting a level of the polypeptide in a sample derived from a host. The 507-bp cDNA coding for the 167-amino-acid, mature form of IL-6SV was isolated and sequenced from a CDNA library derived from activated human \*\*\*macrophages\*\*\*. The examples described the bacterial expression of IL-6SV using the expression vector pQE-60 with a His6 tag and purifit, of the protein from recombinant Escherichia coli on a Ni-Chelate column. Expression via gene therapy is achieved by inserting the cDNA into Moloney murine sarcoma virus-derived vector pMV-7 for the transduction of fibroblasts.

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L12 ANSWER 5 OF 18 CAPLUS COPYRIGHT 2004 ACS on STN
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AN 1996:120813 CAPLUS

DN 124:200000

TI Effects of IL-3 gene-transfected \*\*\*tumor\*\*\* cells on the number and

functions of peritoneal \*\*\*macrophages\*\*\* in vivo

AU Zhang, Weiping; Cao, Xuetao; Ye, Tianxing

CS Department of Immunology, The Second Military Medical University,

Shanghai, 200433, Peop. Rep. China
SO Zhonghua Weishengwuxue He Mianyixue Zazhi (\*\*\*1995\*\*\* ), 15(5), 321-4
CODEN: ZWMZDP; ISSN: 0254-5101

PB Weishenbu Beijing Shengwu Zhipin Yanjiuso

DT Journal

LA Chinese

AB The effects of IL-3 on in vivo secretion by peritoneal \*\*\*macrophages\*\*\* were studied. The nos. of peritoneal \*\*\*macrophages\*\*\* doubled 4 days after inoculation of mice with B16-IL-3 cells, and were increased by 4-5-fold after 10-15 days. The freshly prepd. \*\*\*macrophages\*\*\* from

B16-IL-3 inoculated mice secreted IL-1, IL-6, and TNF, and had a high \*\*\*tumoricidal\*\*\* activity. The cytokine secretion and cytotoxicity were enhanced after induction with LPS in vitro. The la antigen expression was improved. Thus, IL-3 secreted by B16-IL-3 cells in vivo effectively activated the peritoneal \*\*\*macrophages\*\*\*, which may account for the decreased \*\*\*tumorigenicity\*\*\* of the IL-3 gene transfected melanoma cells.

L12 ANSWER 6 OF 18 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on STN

AN 1995:299143 BIOSIS DN PREV199598313443

Adjuvants, endocrines and conserved epitopes: Factors to consider when

designing "therapeutic vaccines". AU Rook, G. A. W.; Stanford, J. L.

Med. Microbiol., UCL Med. Sch., 67-73 Riding House St., London W1P 7LD,

UK SO International Journal of Immunopharmacology, (1995) Vol. 17, No. 2, pp. 91-102

CODEN: IJIMDS. ISSN: 0192-0561.

DT Article

General Review, (Literature Review)

LA English

ED Entered STN: 11 Jul 1995

Last Updated on STN: 11 Jul 1995

AB Research into immunity to complex intracellular parasites has recently Sesearch into immunity to complex intracellular parasites has recently placed emphasis on the identification of peptide sequences recognised by T-cells, often with the dual objectives of finding species-specific protective epitopes, and of understanding selection of Th1 versus Th2 response patterns. In this review it is suggested that although such work is interesting, it will not achieve these objectives, which must, however, be addressed before we can design the new \*\*\*generation\*\*\* of \*\*\*therapeutic\*\*\* vaccines which may eventually replace antimicrobial drugs in the treatment of infection. First, we suggest that the balance of Th1 to Th2 hymphocyte activity is not determined by entitones, but

of Th1 to Th2 lymphocyte activity is not determined by epitopes, but rather by adjuvant effects of microbial components which we have barely begun to define, and local endocrine effects mediated by conversion of

prohormones into active metabolites by enzymes in lymphnode

""macrophages"". Cytokines play a role as mediators within these
pathways. In chronic disease states there is a tendency for T-cell
function to shift towards Th2. We describe immunopathological
consequences of this tendency, including a putative role for agalactosyl IgG, and review evidence for involvement of changes in the endocrine system, brought about not only by the cytokine-hypothalamus-pituitary-adrenal axis, but also by direct actions on peripheral endocrine organs of excess levels of cytokines such as TNF-alpha, TGF-beta and IL-6. We summarise evidence that the epitopes that are targets for protective cell-mediated responses to complex organisms are usually not species specific. In tuberculosis, cellular responses to species-specific components appear to be associated with immunopathology rather than protection. Finally, we discuss how application of these principles has led to remarkable results in the immunotherapy of tuberculosis, including multidrug-resistant disease.

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L12 ANSWER 7 OF 18 CAPLUS COPYRIGHT 2004 ACS on STN
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AN 1995:240035 CAPLUS

DN 122:23868

Therapeutic compositions for use in humans, characterized by a combination of a muramyl peptide and a cytokine

IN Chedid, Louis; Bahr, Georges; Lefrancier, Pierre

Vacsyn S. A., Fr.

SO PCT Int. Appl., 56 pp. CODEN: PIXXD2

DT Patent

LA English

PATENT NO.

KIND DATE WO 9421275

VO 9421275 A1 19940929 WO 1994-FR307 19940321 <--</li>
 W: AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU, JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO,

APPLICATION NO

DATE

19940321

19951113

RU, SD, SE, SK, UA, US, UZ, VN RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

A1 B1 FR 2702659 19940923 FR 1993-3230 19930319 <--FR 2702659 19950825 19941007 FR 1993-3787 19930331 <--FR 2703251 **B3** 19950804 CA 2157758 19940929 CA 1994-2157758 19940321 <--AA AU 9462856 Α1 19941011 19960103 AU 1994-62856 EP 1994-910445 19940321 <--19940321 <--EP 689449 Α1

EP 689449 20021030 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE JP 08511235 T2 19961126 JP 1994-520726 19940321 <--20021115 AT 1994-910445 20030331 PT 1994-910445 AT 226828 PT 689449 E 19940321 19940321

19940321

20030616 ES 1994-910445 19990803 US 1995-522342 ТЗ US 5932208 PRAI FR 1993-3230 Α 19930319 FR 1993-3787 WO 1994-FR307 19930331

w OS MARPAT 122:23868

ES 2187520

AB A therapeutic compn. for use in humans comprises a combination of gtoreq 1 natural or recombinant and preferably human cytokine with .gtoreq.1 muramyl peptide selected from those which, when administered in vivo together with an interferon, also induce an increased in vivo prodn. of an interleukin-1 receptor antagonist, but preferably do not induce any increase in TNF, IL-8 and IL-1 cytokines. The compn. is useful for antiviral and antitumor therapies and/or for promoting restoration of the hematopoietic system, particularly in individuals with a weakened immune system. Studies of the effect of e.g. a mutabutide-interferon combination in an animal toxic shock model are described.

L12 ANSWER 8 OF 18 CAPLUS COPYRIGHT 2004 ACS on STN

AN 1995:210275 CAPLUS

DN 122:532

TI Adenovirus-mediated gene therapy of experimental gliomas AU Perez-Cruet, M. J.; Trask, T. W.; Chen, S-H.; Goodman, J. C.; Woo, S. L. C.; Grossman, R. G.; Shine, H. D.

CS Dep. Neurosurgery, Baylor Coll. Med., Houston, TX, USA SO Journal of Neuroscience Research (\*\*\*1994\*\*\* ), 39(4), 506-11 CODEN: JNREDK; ISSN: 0360-4012

PB Wiley-Liss DT Journal

LA English

The efficacy of adenovirus (ADV)-mediated gene therapy to treat brain \*\*\*tumors\*\*\* was tested in a syngeneic glioma model. \*\*\*Tumor\*\*\* was tested in a syngeneic glorna model. cells were transduced in situ with a replication-defective ADV carrying the herpes simplex virus thymidine kinase (HSV-tk) gene controlled by the Rous sarcoma virus promoter. Expression of the HSV-tk gene enables the transduced cell to convert the drug ganciclovir to a form that is cytotoxic to dividing cells. \*\*\*Tumors\*\*\* were generated in Fischer at a transfer of the caudate nucleus. Eight days later, the \*\*\*tumors\*\*\* were injected either with the ADV carrying the HSV-tk (ADV-tk) gene or a control ADV vector contg. the .beta.-galactosidase (ADV-.beta.gal) gene and the rats were treated with either ganciclovir or saline. \*\*\*Tumor\*\*\* size was measured 20 days after implantation of 9L cells or at death. Rats treated with ADV-.beta.gal and ganciclovir or with ADV-tk and saline had large
\*\*\*tumors\*\*\*\* . No \*\*\*tumors\*\*\*\* were detected in animals treated with ADV-tk and with ganciclovir at doses .gtoreq.80 mg/kg. An infiltrate of \*\*\*macrophages\*\*\* and lymphocytes at the injection site in animals treated with ADV-tk and ganciclovir indicated an active local immune reaction. In survival studies, all animals treated with ADV-tk and ganciclovir have remained alive longer than 80 and up to 120 days after \*\*\*tumor\*\*\* induction whereas all untreated animals died by 22 days.

These results demonstrate that ADV-mediated transfer of HSV-tk to glioma cells in vivo confers sensitivity to ganciclovir, and represents a potential method of treatment of brain \*\*\*tumors\*\*\*.

L12 ANSWER 9 OF 18 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL

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AN 94135569 EMBASE DN 1994135569

IN 1994 (3006)

IT Glycoconjugates as carriers for specific delivery of """therapeutic"" drugs and """genes"".

AU Monsigny M.; Roche A.-C.; Midoux P.; Mayer R.

S Lab. Blochimie Glycoconjugues, Univ. d'Orleans, CNRS, Bat. B, 1 rue Haute, 45071 Orleans Cedex 2, France

Advanced Drug Delivery Reviews, (1994) 14/1 (1-24).

ISSN: 0169-409X CODEN: ADDREP

CY Netherlands

DT Journal; General Review FS 004 Microbiology 016 Cancer

Human Genetics

Immunology, Serology and Transplantation Biophysics, Bioengineering and Medical Instrumentation

030

Drug Literature Index 037

LA English
SL English
AB Cell surface receptors are good candidates to selectively target drugs, oligonucleotides or even genes by making use of their specific ligands. large number of mammalian cells express cell surface sugar-binding proteins, also called 'membrane lectins'. Therefore, sugars may be used as specific recognition signals to specifically deliver biological active components. Tens of membrane lectins with different sugar specificities have been characterized; some of them actively carry their ligands to intracellular compartments, including endosomes, lysosomes and, in some cases, Golgi apparatus. In this review, we summarize the main properties of neoglycoproteins and glycosylated polymers; they have been developed to study the properties of endogenous lectins and to carry various drugs Glycoconjugates have been successfully used to carry biological response onycoconjugates nave been successifully used to carry biological responser modifiers such as N-acety/muramyldipeptide. N-Acety/muramyldipeptide is, in vitro, hundreds of times more efficient in rendering

""macrophages" ""tumoricidal" when it is bound to this type of carrier. In vivo, the N-acety/muramyldipeptide bound to glycoconjugates containing mannose in a terminal nonreducing position, induces the

containing maintose in a criminal indirectoring positivity, indices the eradication of lung metastases, occurring when treatment is started, in 70% of mice; free N-acetylmuramyldipeptide is strictly inactive. Similarly, N-acetylmuramyldipeptide bound to the same glycoconjugates induces an active antiviral effect. Glycoconjugates are also suitable for carrying antisense oligonucleotides specific for viral sequences.

Antisense oligonucleotides protected at both ends and linked through a disulfide bridge to the glycoconjugates are 10 times more efficient than the corresponding free oligonucleotides. Poly-L-lysine containing about 190 lysine residues has been substituted by three components: sugars as recognition signal, antiviral (or antiparasite) agents as therapeutic elements and gluconoic acid as neutralizing and solubilizing agent. This type of neutral, highly water-soluble glycosylated polymer is a very efficient carrier to deliver drugs in infected cells according to the nature of the sugar borne on the polymer and to the specificity of the lectin present at the surface of the infected cells. Finally, poly-L-lysine (190 residues) partially substituted with sugars (60 units) is a polycationic glycosylated polymer which easily makes complexes with plasmids. These complexes are very efficient in transfecting cells in a sugardependent manner. The expression of reporter gene is greatly enhanced when cells are incubated with the plasmid-glycosylated poly-L-lysine complex in the presence of either 100 .mu.M chloroquine or 10 .mu.M fusogenic docosapeptide. Furthermore, this transfection method leads to a much larger number of stable transfectants than the classical method using calcium phosphate precipitate. The general properties of glycosylated proteins and of glycosylated polymers are presented and their efficiency in targeting genes in comparison with that of other available targeted transfection methods is discussed.

L12 ANSWER 10 OF 18 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on

STN

**DUPLICATE 1** 

AN 1993:345126 BIOSIS

DN PREV199396042126

Ex vivo expansion of enriched peripheral blood CD34-positive progenitor cells by stem cell factor, interleukin-1-beta (IL-1-beta), IL-6, IL-3,

interferon-gamma, and erythropoletin.

AU Brugger, Wolfram; Mocklin, Wolfgang; Heimfeld, Shelly; Berenson, Ronald

J.; Mertelsmann, Roland; Kanz, Lothar [Reprint author]

CS Univ. Freiburgh Med. Cent., Dep. Hematology/Oncology, 7800 Freiburg, Germany

SO Blood, (1993) Vol. 81, No. 10, pp. 2579-2584. CODEN: BLOOAW. ISSN: 0006-4971.

DT Article

LA English ED Entered STN: 26 Jul 1993

Last Updated on STN: 27 Jul 1993

AB To provide sufficient numbers of peripheral blood progenitor cells (PBPCs) for repetitive use after high-dose chemotherapy, we investigated the ability of hematopoietic growth factor combinations to expand the number of clonogenic PBPCs ex vivo. Chemotherapy plus granulocyte colony-stimulating factor (G-CSF) mobilized CD34+ cells from 18 patients with metastatic solid \*\*\*tumors\*\*\* or refractory lymphomas were cultured for up to 28 days in a liquid culture system. The effects of interleukin-1-beta (IL-1), IL-3, IL-6, granulocyte-macrophage-CSF (GM-CSF), G-CSF, macrophage-CSF (M-CSF), stem cell factor (SCF), erythropoietin (EPO), leukemia inhibitory factor (LIF), and interferon-gamma, as well as 36 combinations of these factors were tested. A combination of five hematopoietic growth factors, including SCF, EPO, IL-1, IL-3, and IL-6, was identified as the optimal combination of growth factors for both the expansion of total nucleated cells as well as the expansion of clonogenic progenitor cells. Proliferation peaked at days 12 to 14, with a median 190-fold increase (range, 46- to 930-fold) of total clonogenic progenitor cells. Expanded progenitor cells generated myeloid (colony-forming unit-granulocyte-macrophage), erythroid (burst-forming unit-erythroid), as well as multilineage (colony-forming unit-granulocyte, erythrocyte, \*\*\*monocyte\*\*\*, megakaryocyte) colony-forming units. The number of multilineage colonies increased 250-fold (range, 33- to 589-fold) as compared with pre-expansion values. Moreover, the absolute number of early hematopoietic progenitor cells (CD34+/HLA-DR-; CD34+/CD38-), as well as the number of 4-HC-resistant progenitors within expanded cells increased significantly. Interferon-gamma was shown to synergize with the 5-factor combination, whereas the addition of GM-CSF significantly decreased the number of total clonogenic progenitor cells. Large-scale expansion of PB CD34+ cells (starting cell number, 1.5 times 10-6, CD34+ cells) in autologous plasma supplemented with the same 5-factor combination resulted in an equivalent expansion of progenitor cells as compared with the microculture system. In summary, our data indicate that chemotherapy plus G-CSF-mobilized PBPCs from \*\*\*cancer\*\*\* patients can be effectively expanded ex vivo. Moreover, our data suggest the feasibility of large-scale expansion of PBPCs, starting from small numbers of PB CD34+ cells. The number of cells expanded ex vivo might be sufficient for repetitive use after high-dose chemotherapy and might be candidate cells for \*\*\*therapeutic\*\*\* \*\*\*\*gene\*\*\* transfer.

L12 ANSWER 11 OF 18 BIOSIS COPYRIGHT (c) 2004 The Thomson Corporation. on

STN

**DUPLICATE 2** 

AN 1993:522399 BIOSIS DN PREV199396135806

Efficient adenovirus-mediated gene transfer into human blood

\*\*\*monocyte\*\*\* - derived \*\*\*\*macrophages\*\*\* .

AU Haddada, Hedi [Reprint author]; Lopez, Manuel; Martinache, Chantal; Ragot, Thierry; Abina, Mohammed Amine; Perricaudet, Michel

CS CNRS UA 1301, Inst. Gustave Roussy, PR2 39 rue Camille Desmoulins,

SO Biochemical and Biophysical Research Communications, (1993) Vol. 195, No. 3, pp. 1174-1183.

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ACTIVITY PRODUCTION IN-VITRO BY EQUINE PERITONEAL
      CODEN: BBRCA9, ISSN: 0006-291X
DT Article
LA English
ED Entered STN: 19 Nov 1993
                                                                                                                                                                                                        ***MACROPHAGES*** .

AU MORRIS D D [Reprint author]; MOORE J N; CROWE N; FISCHER J K
CS DEP LARGE ANIMAL MED, COLL VET MED, UNIV GA, ATHENS, GA 30602,
      Last Updated on STN: 19 Nov 1993
                                                                                                                                                                                                        SO Cornell Veterinarian, (1991) Vol. 81, No. 3, pp. 267-276.
CODEN: COVEAZ. ISSN: 0010-8901.
AB The efficiency of gene transfer into human blood ***monocyte -derived ***macrophages*** has been evaluated using a
     replication-defective adenovirus vector harboring a lac Z gene of E. coli as a reporter gene. Whereas, no beta-galactosidase activity was found in freshly infected purified ***monocytes***, 40% to 80% of infected ***macrophages*** which derived from these ***monocytes*** showed a beta-galactosidase activity, 2 to 4 days after infection and lasted for at
                                                                                                                                                                                                        DT Article
                                                                                                                                                                                                        FS
                                                                                                                                                                                                                 BA
                                                                                                                                                                                                       LA ENGLISH
ED Entered STN: 13 Aug 1991
Last Updated on STN: 13 Aug 1991
                                                                                                                                                                                                       AB This study evaluated the effect of dexamethasone on endotoxin-induced production of ***tumor*** necrosis factor (TNF) activity in vitro by equine peritoneal ***macrophages***. Peritoneal ***macrophages*** from adult horses were cultured in the presence of dexamethasone (1-100 ...mu.M) for various time periods (2 hour, 0.5 hour, 0 hour) prior to the
      beta-galactosiase activity, 26 4 days after infection and lasted for at least 3 weeks. Moreover, beta-galactosidase activity was found in infected ***monocyte*** / ***macrophages*** 7 days after their injection into a human ***tumor*** preestablished in nude mice. These data indicate that it is possible to transfer and stably express a ****gene*** of potential ***therapeutical*** function into human ***monocyte*** -derived ***macrophages*** using an adenovirus vector.
                                                                                                                                                                                                              addition of endotoxin (5 ng/ml), then the secretion of TNF activity was evaluated. Macrophage supernatant concentrations of TNF activity were
                                                                                                                                                                                                              estimated by a modified in vitro cytotoxicity bioassay using the murine fibrosarcoma cell line, WEHI 164 clone 13. An experiment was performed to determine whether dexamethasone interfered with the cytolytic bioassay's
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      on STN
AN 93074734 EMBASE
DN 1993074734
TI Blood transfusion related adult respiratory distress syndrome.
                                                                                                                                                                                                              ability to detect TNF activity. The endotoxin-induced TNF activity production by equine peritoneal ****macrophages*** was significantly reduced by co-incubation with 100 .mu.M dexamethasone, but not by tested
                                                                                                                                                                                                              concentrations of dexamethasone less than 100 .mu.M. This concentration of dexamethasone greatly exceeds those ***generally*** attained by ***therapeutic*** use of dexamethasone in horses. Preincubation time
 AU Malouf M.; Glanville A.R.

    So Division of Respiratory Medicine, Concord Hospital, Concord, NSW, Australia Anaesthesia and Intensive Care, (1993) 21/1 (44-49).
                                                                                                                                                                                                              did not affect the ability of 100 .mu.M dexamethasone to reduce TNF production by equine ***macrophages***. The quantitation of equine TNF activity by its cytolytic bioassay was not altered by dexamethasone
      ISSN: 0310-057X CODEN: AINCBS
 CY Australia
         Journal; General Review
FS 005 General Pathology and Pathological Anatomy
015 Chest Diseases, Thoracic Surgery and Tuberculosis
                                                                                                                                                                                                        L12 ANSWER 15 OF 18 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL
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                    Hematology
Drug Literature Index
      025
                                                                                                                                                                                                              on STN
      037
                                                                                                                                                                                                         AN 90308336 EMBASE
                                                                                                                                                                                                        DN 1990308336
LA English
SL English
AB Adult respiratory distress syndrome (ARDS) is a rare but important
                                                                                                                                                                                                        TI Cellular properties of ***cancer*** .
AU Miller F.R.
      complication of blood transfusion because it has a mortality rate of 50-60%. ARDS is characterised by noncardiogenic pulmonary oedema and is often associated with major trauma and/or sepsis. Clinical features include dyspnoea, tachypnoea, chills and extensive crepitations. The pathogenesis has not been elucidated completely and a number of hypotheses
                                                                                                                                                                                                              Michigan Cancer Foundation, Detroit, MI, United States
Current Opinion in Oncology, (1990) 2/1 (152-156).
ISSN: 1040-8746 CODEN: CUOOE8
                                                                                                                                                                                                        CY United States
                                                                                                                                                                                                                 Journal; General Review
      have been proposed. Factors which have been implicated include neutrophil sequestration and complement activation, ***macrophages***, metabolites of the arachidonic acid cascade and cytokines, all of which
                                                                                                                                                                                                        FS 016 Cancer
                                                                                                                                                                                                       LA English
                                                                                                                                                                                                                 English
      contribute to the amplification of the inflammatory process. In particular, leucoagglutinins have been implicated with blood transfusions.

Treatment is ***generally*** supportive as specific
***therapeutic*** strategies remain largely unproven.
                                                                                                                                                                                                        AB It is generally agreed that spontaneously developing ***cancers*** are usually of monoclonal origin and grow autonomously. However,
                                                                                                                                                                                                             usually of monoclonal origin and grow autonomously. However,
"""Cancers" are not packets of identical cells growing uncontrollably.
""Cancers" contain many subpopulations of neoplastic cells that
differ in many clinically relevant characteristics such as growth rate,
ability to metastasize, response to """therapeutic"" modalities, and
""genetic"" stability. In addition to this 'donal heterogeneity,'
normal cells of the tissue or origin (or site of metastasis) and
infiltrating host cells (lymphocytes, ""macrophages", leukocytes)
are present as well as extracellular matrix components. Depending on the
vascularization of the ""tumor", pH gradients and relative hypoxia
generate an additional level of heterogeneity. All of these factors are
interactive, creating a dynamic system that is able to evolve or progress
under the selective pressures of the host (homeostatic mechanisms and
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      on STN
 AN 92344033 EMBASE
DN 1992344033
 TI Congenital monoblastic leukemia and 9;11 translocation. A case report.
AU Monpoux F.; Sirvent N.; Sudaka I.; Mariani R. CS Clinique Medicale Infantile, Hopital de Cimiez, Av. Victoria,06003 Nice,
France
SO Pediatrie, (1992) 47/10 (691-694).
ISSN: 0031-4021 CODEN: PEDRAN
                                                                                                                                                                                                              under the selective pressures of the host (homeostatic mechanisms and
                                                                                                                                                                                                              immune responses) or of therapeutic interventions.
                                                                                                                                                                                                       L12 ANSWER 16 OF 18 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
CY France
DT Journal, Article
      007 Pediatrics and Pediatric Surgery
016 Cancer
025 Hematology
                                                                                                                                                                                                        on STN
AN 90062241 EMBASE
 FS 007
                                                                                                                                                                                                        DN 1990062241

TI Acute monoblastic leukemia: a unique subtype - a review from the Childrens
***Cancer*** Study Group.

AU Odom L.F.; Lampkin B.C.; Tannous R.; Buckley J.D.; Hammond G.D.
 LA French
 SL French: English
 AB Acute leukemia in the newborn child is a rare event. The clinical and
                                                                                                                                                                                                       CS Children's Hospital of Denver, University of Colorado Health Sciences Center, Denver, CO, United States SO Leukemia Research, (1990) 14/1 (1-10). ISSN: 0145-2126 CODEN: LEREDD
      biological characteristics differ from those normally encountered in the older child. ***Tumoral*** syndrome and extra-medullar locations are frequently described in the literature. Many authors have noted the
     difficulty of diagnosis due to the immaturity of the malignant proliferation. While it is ""generally" agreed that ""therapeutic" abstention is justified in the leukemoid reaction in Down's syndrome, the choice is debatable in the phenotypically intact newborn. For this reason, blastic caryotype analysis is essential and may
                                                                                                                                                                                                                 United Kingdom
                                                                                                                                                                                                       DT Journal; General Review
FS 005 General Pathology and Pathological Anatomy
                                                                                                                                                                                                                            Internal Medicine
                                                                                                                                                                                                              006
      provide guidelines when considering treatment. We report on a case history of acute monoblastic leukemia with translocation 9;11 that was diagnosed
                                                                                                                                                                                                                            Neurology and Neurosurgery
                                                                                                                                                                                                               800
                                                                                                                                                                                                              016
                                                                                                                                                                                                                            Cancer
      at birth in a normal newborn infant. The juxtaposition of c-ets 1 protooncogene and the beta-interferon gene has been associated with this kind of cytogenetic disease and probably constitutes a model for human
                                                                                                                                                                                                              025
                                                                                                                                                                                                                         Hematology
                                                                                                                                                                                                        LA English
                                                                                                                                                                                                        SL English
                                                                                                                                                                                                        AB The acute non-lymphocytic leukemias (ANLL) are generally treated as a homogeneous group. However, the literature is replete with articles alluding to distinctive features of acute monoblastic leukemia (AMoL).
 L12 ANSWER 14 OF 18 BIOSIS COPYRIGHT (c) 2004 The Thomson
                                                                                                                                                                                                              This review addresses the unique clinical, laboratory, epidemiological, and therapeutic features of AMoL. Leukemic monoblasts are distinguished
Corporation, on
                                                                              DUPLICATE 3
                                                                                                                                                                                                              from other cells in the myelocytic series by physical properties such as greater adhesiveness, deformability, and motility. Patients with AMoL often exhibit hyperleukocytosis, disseminated intravascular coagulation,
AN 1991:365871 BIOSIS
DN PREV199192054096; BA92:54096
TI DEXAMETHASONE REDUCES ENDOTOXIN-INDUCED ***TUMOR***
NECROSIS FACTOR
                                                                                                                                                                                                              and extramedullary involvement, particularly in the skin, gingiva, and
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central nervous system (CNS). AMoL occurs predominantly in adults over 40 and children under 10, fifty percent of whom are under 2 years of age at diagnosis. Its relatively common occurrence in infants parallels the high rate of proliferation of \*\*\*monocytes\*\*\* in that age group. Additionally, its occurrence in young children appears to be associated with in utero exposure to marijuana and parental exposure to pesticides and solvents. \*\*\*Therapeutic\*\*\* results are \*\*\*generally\*\*\* poor due to high rates of fatal complications during induction, induction failures, and frequent extramedullarly and medullary relapses. This poor outcome is particulary noted in infants. Higher remission induction rates attained with epipodophyllotoxins and incorporation of bone marrow transplantation have not yet resulted in substantial improvement of long-term outcome. Recurrence of disease in the CNS is minimized by the use of intensive CNS presymptomatic treatment, usually incorporating irradiation. Our review suggests that unique and innovative treatment strategies are needed to improve outcome for patients with AMoL.

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AN 1987:82084 BIOSIS DN PREV198783040662: BA83:40662

TI BIOLOGICAL PROPERTIES AND MOLECULAR BIOLOGY OF THE HUMAN MACROPHAGE GROWTH
FACTOR COLONY STIMULATING FACTOR 1.

AU RALPH P [Reprint author]; WARREN M K; NAKOINZ I; LEE M-T; BRINKLEY

SAMPSON-JOHANNES A; KAWASAKI E S; LADNER M B; STRICKLER J E;

CS CETUS CORP, 1400 FIFTY-THIRD ST, EMERYVILLE, CALIF 94608, USA SO Immunobiology, (1986) Vol. 172, No. 3-5, pp. 194-204. CODEN: IMMND4. ISSN: 0171-2985.

DT Article FS BA

ENGLISH

ED Entered STN: 7 Feb 1987

Last Updated on STN: 7 Feb 1987

AB CSF-1 is a growth and differentiation factor for the production of mononuclear \*\*\*phagocytes\*\*\* from undifferentiated bone marrow progenitors. In addition to previously described effects on mature cells, we show here that CSF-1 stimulates the production by \*\*\*monocytes\*\* of interferon, \*\*\*tumor\*\*\* necrosis factor, and myeloid CSF that produces mainly mixed neutrophil-macrophage colonies in bone marrow culture. Pretreatment with CSF-1 also promotes resistance to viral

infection and """tumor"" cytotoxicity in murine peritoneal
""macrophages"". Based on amino acid sequence data of purified human
urinary and murine L cell CSF-1, we have cloned the complementary DNA
(cDNA) from messenger RNA (mRNA) of the human CSF-1 producing MIA PaĈa

cell line. The cDNA species a 32 amino acid signal peptide followed by a protein of 224 amino acids. Several facts suggest, however, that one-third of the molecule at the C-terminal end is processed off intracellularly to derive the secreted growth factor. The gene is about 18 kilobases (kb) in length and contains 9 exons. Although here appears to be a single copy gene for CSF-1, cells expressing the factor contain several mRNA species, suggesting that the gene may have several functions or levels of regulation. High level expression of the recombinant protein will alllow preclinical testing in several disease models for therapeutic efficacy that has been suggested that in vitro and in vivo biological properties of CSF-1.

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on STN

AN 84080135 EMBASE

DN 1984080135

TI Immunostimulation, Clinical and experimental perspectives

AU Drews J.

CS Sandoz AG, Pharma Forschung und Entwicklung, CH-4002 Basel, Switzerland

SO Klinische Wochenschrift, (1984) 62/6 (254-264).

CODEN: KLWOAZ CY Germany

DT Journal

FS 037 Drug Literature Index 026 Immunology, Serology and Transplantation

AB Three classes of immunostimulating drugs are described, each representing a different approach to the problem of pharmacological immunostimulation. The rationale for the use of microbes or microbial agents as immunostimulators rests on the fact that some micro-organisms, especially those that replicate intracellularly, carry a special potential to activate \*\*\*macrophages\*\*\*. Clinically, the use of these agents in patients with \*\*\*tumors\*\*\* and infections has been disappointing; however, there have been positive exceptions like the responsiveness of melanomas and bladder carcinomas to the injection of BCG. Many of the inconclusive results may be due to insecurities in the dosage of microbial preparations and to a general lack in standardization. Some structures with high efficacy and low toxicity which have recently evolved from this field deserve further investigation. A number of structurally unrelated synthetic compounds was found to influence immune parameters. Levamisole can today be classified as an immunostimulating drug with limited utility in recurring infections and in chronic polyarthritis. Several

immunostimulating drugs which have attracted interest contain a purine as the effective component. This is not surprising in view of the fact that many genetically determined immunodeficiencies can be traced to defe enzymes which play a crucial role in purine biosynthesis. Finally, the potential role of lymphokines as stimulators of the immunosystem is briefly described. Some of these glycoproteins have recently become available for clinical trials. Others will be made available through

\*\*\*genetic\*\*\* engineering. The \*\*\*therapeutic\*\*\* utility of these
compounds is not yet clear; they will, however, be of great value as probes for the study of immune functions and for the development of immunopharmacology.

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